In re: Application of TERADA et al.

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## **AMENDMENTS TO THE CLAIMS**

This listing of claims will replace all prior versions, and listings, of claims in the application:

## **Listing of Claims:**

- (Currently amended). A method for identifying a drug candidate for promoting tissue-specific differentiation of an embryonic stem cell (ES), the method comprising the steps of:
  - (A) providing a library of test substances, the library comprising at least a first test substance and a second test substance, the first and second test substances having different molecular structures;
  - (B) providing an *in vitro* culture of embryonic stem cells (ES) cultured in hanging drops for 2 days to produce a culture of embryoid bodies, the culture being divided into at least a first subculture and a second subculture and;
  - (C) culturing the first and second subcultures for at least about 5 days in the absence of a test substance on a collagen coated culture plate coated with collagen or without collagen;
  - (D) contacting the first subculture with a first test substance and a second subculture with a second another test substance from the library of test substances;
  - (E) culturing the first and second subcultures for 7 to 18 days; and
  - (F) analyzing the cells in the first and second subcultures for increased tissuespecific gene expression.
- 2. (Cancel).
- 3. (Previously presented) The method of claim 1, wherein the embryonic stem cells are mammalian embryonic stems cells.

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## 4. (Cancel)

- 5. (Previously presented) The method of claim 1, wherein the embryonic stem cells are murine R1 embryonic stem cells.
- 6. (Previously presented) The method of claim 1, wherein the embryonic stem cells are human embryonic stem cells.

## Claims 7-13. (Cancelled)

- 14. (Previously Presented) The method of claim 1, wherein the step (F) of analyzing the cells in the first and second subcultures for increased tissue-specific gene expression comprises isolating mRNA from the first and second subcultures.
- 15. (Original) The method of claim 14, wherein total cellular RNA is isolated from the first and second subcultures.
- 16. (Previously Presented) The method of claim 14, wherein the step (F) further comprises reverse-transcribing the mRNA to create cDNA.
- 17. (Previously Presented) The method of claim 1, wherein the step (F) of analyzing the cells in the first and second subcultures for increased tissue-specific gene expression comprises performing a polymerase chain reaction (PCR).
- 18. (Original) The method of claim 14, wherein the isolated mRNA is immobilized on a substrate.

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19. (Original) The method of claim 18, wherein the substrate is contacted with a probe that specifically hybridizes to the tissue-specific mRNA.

20. (Previously Presented) The method of claim 1, wherein the step (F) of analyzing the cells in the first and second subcultures for increased tissue-specific gene expression is performed using gene chip technology.